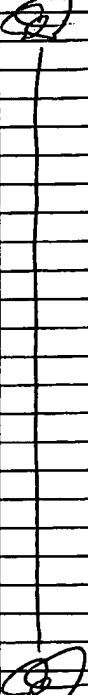




PTO/SB/08A (08-03)

Substitute for form 1449A/PTO			Complete if Known		
INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)			Application Number	10/669,099	
			Filing Date	September 22, 2003	
			First Named Inventor	Gorfine, Stephen R.	
			Art Unit	1654	
			Examiner Name	Tate, Christopher Robin	
Sheet	1	of	7	Attorney Docket Number	010692-004532US

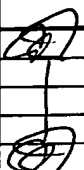
U.S. PATENT DOCUMENTS+						
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		Number	Kind Code ² (if known)			
	1	US-814,408		03-06-1906	Steele	
	2	US-2,840,080		06-24-1958	Clark	
	3	US-3,419,571		12-31-1968	Di Carlo et al.	
	4	US-4,118,480		10-03-1978	Williams	
	5	US-4,226,849		10-07-1980	Schor	
	6	US-4,292,299		09-29-1981	Suzuki et al.	
	7	US-4,371,516		02-01-1983	Gregory et al.	
	8	US-4,514,384		04-30-1985	Gallina	
	9	US-4,608,249		08-26-1986	Otsuka et al.	
	10	US-4,654,209		03-31-1987	Leslie et al.	
	11	US-4,657,906		04-14-1987	Emmet et al.	
	12	US-4,683,242		07-28-1987	Poser	
	13	US-4,746,675		05-24-1988	Makino et al.	
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	16	US-5,059,603		10-22-1991	Rubin	
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	20	US-5,504,117		04-02-1996	Gorfine	
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	22	US-5,595,753		01-27-1997	Hechtman	
	23	US-5,595,970		01-21-1997	Garfield et al.	
	24	US-5,693,676		12-07-1997	Gorfine	

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Examiner Initials*	Cite No. ¹	Foreign Patent Document		Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T ⁶
		Country Code ³	Number ⁴ (Kind Code ² (if known))				
	25	DE	304,730	03-01-1989	Dunendorfer et al.		<input type="checkbox"/>
	26	AU	74545/94	08-19-1994	Koren Laboratories Pty., Ltd.		<input type="checkbox"/>
	27	AU	PM 0971	03-22-1995	Koren Laboratories Pty., Ltd.		<input type="checkbox"/>
	28	AU	PM 4247	03-22-1995	Koren Laboratories Pty., Ltd.		<input type="checkbox"/>
	29	AU	PM 6395	07-14-1994	Koren Laboratories Pty., Ltd.		<input type="checkbox"/>
	30	DE	4,038,203	06-04-1992	Bonnacker et al.		<input type="checkbox"/>
	31	JP	62-67017	03-26-1987	Sawai et al.		<input type="checkbox"/>
	32	WO	98/36733	08-27-1998	Kamm, Michael Albert et al.		<input type="checkbox"/>
	Examiner Signature: <i>C. R. Tate</i>						Date Considered: <i>12/12/05</i>

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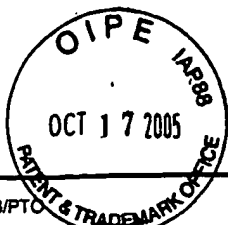
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Substitute for form 1449A/PTO INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)			Complete if Known		
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			Filing Date	September 22, 2003	
			First Named Inventor	Gorfine, Stephen R.	
			Art Unit	1654	
Examiner Name	Tate, Christopher Robin				
Sheet	2	of	7	Attorney Docket Number	010692-004532US

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		Number ⁴						
		Country Code ³	Kind Code ⁵ (if known)					
	33	WO	95/06466	A1	03-09-1995	Koren Laboratories Pty., Ltd.		
	34	ZA	77/01135		02-25-1977	Prinsloo, J.		
	35	JP	04-091036		03-24-1996	Hashioka Tatsuya		
	36	WO	87/03490		06-18-1987	Key Pharmaceuticals, Inc.		
	37	WO	98/37886		09-03-1998	Antropoli		

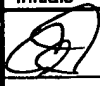
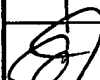
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PTO/SB/08B (08-03)

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			First Named Inventor	Gorfine, Stephen R.	
			Art Unit	1654	
			Examiner Name	Tate, Christopher Robin	
Sheet	3	of	87	Attorney Docket Number	010692-004532US

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	38	Ahner et al., "Organic Nitrate Esters: Clinical Use and Mechanisms of Actions." <i>Pharmacol. Reviews</i> 43(3):351-353 (1991).	
	39	Budavari et al. (Eds.), "The Merck Index." pp. 198 and 821 (isopropyl nitrate); 11th Edition, Merck & Co., Rahway, N.J. (1989)	
	40	Bult et al., "Nitric oxide as an inhibitory non-adrenergic non-cholinergic neurotransmitter." <i>Nature</i> , 345:346-347, 1990.	
	41	Burleigh, D. and D'Mello, A., "Physiology and Pharmacology of the Internal Anal Sphincter." Chapter 2, pages 22-41, in <i>Coloproctology and The Pelvic Floor</i> , Henry and Swash (Eds.), Butterworth Publishers, Stoneham, Mass., USA and London, England, ISBN 0-407-00352-5 (1985)	
	42	Burleigh, D., "Ng-Nitro-L-Arginine Reduces Nonadrenergic Relaxations of Human Gut." <i>Gastroenterology</i> , 102: 679-683 (1992)	
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	44	Enck et al., "Spontaneous variation of anal 'resting' pressure in healthy humans." <i>Am. J. Physiol.</i> , 261(5 Pt 1):G823-G826 (1991)	
	45	Farouk et al., "Sustained Internal Sphincter Hypertonia in Patients with Chronic Anal Fissure." <i>Dis. Col. Rect.</i> , 37:424-429 (1994)	
	46	Feelisch et al., "Correlation between nitric oxide formation during degradation of organic nitrates and activation of guanylate cyclase." <i>Eur. J. Pharmacol.</i> , 139:19-30 (1987)	
	47	Fung et al., "Biochemical mechanism of organic nitrate action." <i>Am. J. Cardiol.</i> , 70:48-10B (1992)	
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	52	Guillemot et al., "Action of in situ nitroglycerin on upper anal canal pressure of patients with terminal constipation. A pilot study." <i>Dis. Col. Rect.</i> , 36(4):372-376 (1993)	

Examiner Signature	C. R. Tate	Date Considered	12/12/05
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			Filing Date	September 22, 2003		
			First Named Inventor	Gorfine, Stephen R.		
			Art Unit	1654		
			Examiner Name	Tate, Christopher Robin		
Sheet	4	of	8	7	Attorney Docket Number	010692-004532US

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	53	Huff et al., (Eds.), "Physicians' Desk Reference." 41st Edition, Medical Economics Company, Oradell, N.J., at pp. 780, 1176-1178, 1533 and 1984-85 (1987)	
	54	Ignarro et al., "Nitric oxide and cyclic GMP formation upon electrical field stimulation cause relaxation of corpus cavernosum smooth muscle." <i>Biochem. Biophys. Res. Commun.</i> , 170:843-850, 1990.	
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	57	Keighley, R. and Williams, N., "Anatomy and Physiology." Chapter One, in <u>Surgery of the Anus, Rectum and Colon</u> , London, W.B. Saunders (1993)	
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	68	Mims Annual Drug Reference, Fifteenth Edition, 2-75, page 2042 (1991)	

Examiner Signature	C. R. Tate	Date Considered	12/12/05
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			First Named Inventor	Gorfine, Stephen R.	
			Art Unit	1654	
			Examiner Name	Tate, Christopher Robin	
Sheet	5	of	87	Attorney Docket Number	010692-004532US

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Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
	69	O'Kelley, T. et al, "Nerve mediated relaxation of the human internal anal sphincter: the role of nitric oxide." <i>Gut</i> , 34: 689-693 (1993)	
	70	Rattan, S. and Chakder, S., "Role of nitric oxide as a mediator of internal anal sphincter relaxation." <i>Am. J. Physiol.</i> G107-G112 (1992)	
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	74	Speakman, C. et al., "Sphincter injury after anal dilation demonstrated by anal endosonography." <i>Br. J. Surg.</i> , 78: 1429-1430 (December 1991)	
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	82	Dodi, "Hot or cold in anal pain? A study of the changes in internal anal spinchter pressure profiles." <i>Dis. Col. Rect.</i> , 29(4):248-51 (1986)	
	83	Farid, M. "Internal anal sphincter relaxation with topical nitroglycerin in the treatment of some benign anal conditions." <i>Br. J. Surg.</i> , 84:1 (1997)	
	84	Farouk, R. et.al. "Changing patterns of treatment for chronic anal fissure." <i>Annual Review College Surgeons England</i> ; 80:194-6 (1998)	

Examiner Signature	C. L. Tate	Date Considered	12/12/05
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* EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 809. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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			Filing Date	September 22, 2003	
			First Named Inventor	Gorfine, Stephen R.	
			Art Unit	1654	
			Examiner Name	Tate, Christopher Robin	
Sheet	6	of	87	Attorney Docket Number	010692-004532US

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	85	"Glyceryl trinitrate for anal fissure?" <i>Drug and Therapeutics Bulletin</i> ; 36:55-6 (1998)	
	86	Gorfine, S.R. "Topical nitroglycerin ointment for anal fissures and ulcers." <i>New England Journal of Medicine</i> ; 333: 1156-7 (1995)	
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	91	Lowenstein, B. et.al. Treatment of proctalgia fugax with topical nitroglycerin: report of a case. <i>Dis. Col. Rect.</i> ; 41:667-8. (1998)	
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	93	Lysy, J. et.al. "Treatment of chronic anal fissure with isosorbide dinitrate: long-term results and dose determination." <i>Dis. Col. Rect.</i> , 41:1406-10 (1998)	
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	98	Pitt, J. et.al. "Chemical sphincterotomy for anal fissure." <i>Colorectal disease</i> 1998; 2-8.	
	99	Pitt, "Double-blind, randomized, placebo-controlled trial of oral indoramin to treat chronic anal fissure." ASCRS Meeting, June 2000, Poster Presentation P2 (2000)	
	100	Romano, "A critical appraisal of pathogenesis and morbidity of surgical treatment of chronic anal fissure." <i>J. Am. Coll. Surg.</i> , 178:600-4 (1994)	

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Sheet	7	of	87	Attorney Docket Number	010692-004532US

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	101	Schouten, "Relationship between anal pressure and anodermal blood flow: The vascular pathogenesis of anal fissures." <i>Dis. Col. Rect.</i> , 37(7):664-9 (1994)	
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